

IV

CELLULAR PROTEINS

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LOCUS p53-php53c1 1317 bp ss-mRNA PRI 22-OCT-1992
 DEFINITION Human p53 cellular tumor antigen mRNA, complete cds.
 ACCESSION X02469 M60950
 KEYWORDS antigen; tumor antigen.
 SOURCE Human, cDNA to mRNA, clone php53c1 from a cDNA library of SV40 transformed fibroblasts.
 REFERENCE 1 (bases 1 to 1317)
 AUTHORS Zakut-Houri,R., Bienz-Tadmor,B., Givol,D. and Oren,M.
 TITLE Human p53 cellular tumor antigen: cDNA sequence and expression in COS cells
 JOURNAL EMBO J. 4, 1251-1255 (1985)
 REFERENCE 2 (bases 1 to 1317)
 AUTHORS Rideout,W.M.III., Coetzee,G.A., Olumi,A.F. and Jones,P.A.
 TITLE 5-methylcytosine as an endogenous mutagen in the human ldl receptor and p53 genes
 JOURNAL Science 249, 1288-1290 (1990)
 REFERENCE 3 (M14694 and M14695 shown in alignment below)
 AUTHORS Harris,N., Brill,E., Shohat,O., Prokocimer,M., Wolf,D., Arai,N. and Rotter,V.
 TITLE Molecular basis for heterogeneity of the human p53 protein
 JOURNAL Mol. Cell. Biol. 6, 4650-4656 (1986)
 REFERENCE 4 (K031991 shown in alignment below)
 AUTHORS Harlow,E., Williamson,N.M., Ralston,R., Helfman,D.M. and Adams,T.E.
 TITLE Molecular cloning and in vitro expression of a cDNA clone for human cellular tumor antigen p53
 JOURNAL Mol. Cell. Biol. 5, 1601-1610 (1985)
 COMMENT Clone php53c1 is the wild-type form of p53, isolated from an SV40 transformed fibroblast cell line. It was first thought that p53, a cell-cycle regulatory protein, was an oncogene. This has since been disproven, for it has been shown that the clones that were first isolated and tested were mutants. In fact, the p53 gene is a tumor suppressor. The mutated forms are thought to be trans-dominant over the wild-type p53. The mutations cause a conformational change in the protein which facilitates its binding to the heat shock protein hsp70. The two forms of p53 and hsp70 become bound together in long-lived unproductive complexes. Since these complexes aggregate in the cytoplasm, the nucleus is deprived of p53. With the negative regulatory effect of p53 reduced or eliminated, the cell is more easily able to proliferate. However, a different mechanism of inactivation is employed by the viral oncoproteins: the large T-antigen of SV40 and the E1B 55kD protein of adenovirus. Instead of the hsp70, mutant p53 complex trapping the wild-type p53 in inactive structures it is the viral oncoproteins that sequester the wild type p53 and inactivate it. Human papillomavirus uses yet another mechanism. The E6 protein of the oncogenic HPV types bind wild-type p53 and stimulate its destruction; although at this point there is no evidence that E6 proteins of noncogenic HPV types, such as HPV-6 or HPV-11, can associate with wild-type p53 in vivo (Werness et al. Science 248, 76-79, Scheffner et al. Cell 63, 1129-1136). It is however possible that the binding assay used to determine this fact was not sensitive enough to detect low affinity binding. The degradation of p53 by HPV E6 is ATP dependant and the ubiquitin-dependant protease system has been suggested to be involved. Since the large T antigen of SV40 also complexes with wild-type p53 and does not target it for degradation, there must be an additional signaling mechanism not yet elucidated involved in the interaction with the HPV E6 protein.
 The p53 wild type php53c1 coding region and four variants are shown in the alignment below. The p53 coding regions which are represented by the accession numbers M14694 and M14695 have both been isolated from the human transformed cell line SV-80. The two forms differ from one another by a single base pair substitution and consequently accounts for the change in electrophoretic mobility of the two proteins. Harris et al. believe that this heterogeneity is due to gene polymorphism.

The coding region represented by accession number K03199 has been isolated from the human vulva carcinoma cell line A431.

BASE COUNT 295 a 408 c 352 g 262 t
 ORIGIN 2 bp upstream of XbaI site; chromosome 17p13.

```

  1 gtctagagcc accgtccagg gaggcaggtag ctgctggct ccggggacac tttgcgttcg
  61 ggctgggagc gtgttcca cgacggtag acgttccct ggattggcag ccagactgcc
 121 ttccgggtca ctgccATGga ggagccgcag tcagatcta gcgtcgagcc ccctctgagt
      p53 start ->
181 cagggaaacat ttccagacat atggaaaacta cttcctgaaa acaacgttct gtcccccttg
241 ccgtcccaag caatggatga tttgatgtg tccccggacg atattgaaca atggttcaact
301 gaagaccagg gtccagatga agctcccaaga atgcccagg ctgctcccc cgtGGCCCT
      ->
361 GCACcagcac ctccatacc ggcGGCCCT GCACcagccc cctccgtggcc cctgtcatct
      <- direct rpt      ->      <- direct rpt
421 tctgtccctt cccagaaaac ctaccaggc agctacgggt tccgtctggg ctcttgcatt
481 tctggacag ccaagtctgt gacttgcacg tactccctg ccctaacaat gatgtttgc
541 caactggcca agacctgccc tttgcagctg tgggttattt ccacaccccc gcccggcacc
601 cgcgtccgcg ccatggccat ctacaaggcag tcacagcaca tgacggaggt tttggggcgc
661 tggcccccacc atgagcgctg ctccatggc gatggctgg cccctctca gcatcttata
721 cgagtggaaag gaaatttgcg tttggatgtt ttggatgaca gaaacactt tcgatcatgt
781 gtgggtgtgc cctatggccg gcctgggtt ggctctgact gtaccacccat ccactacaac
841 tacatgtta acagttccctg catggccgc atgaaccggg gggccatcc caccatcatc
901 acactggaaag actccagggtt taatctactg ggacggaaaca gctttgggt gcgtgtttgt
961 gcctgtccctg ggagagaccc ggcacacccgg gaagagaatc tccgcaagaa aaaaaacccct
1021 caccacggc tgcccccagg gggccactaag cgacactgc ccaacaacac cagctccctt
1081 ccccaagccaa agaagaaaacc actggatggaa gaatatttca cccttcagat ccgtgggggt
1141 gagcgcctcg agatgttccg agagctgaat gggccctgg aactcaagga tgcccaaggct
1201 gggaaaggagc caggggggag cagggtcacttccagccacc tgaagtccaa aaagggtcag
1261 tctacctccc ggcataaaaaa actcatgttc aagacagaag ggcctgactc agacTGA
      <- p53 end
  //
```

P53.WTPHP53C1	ATGGAGGAGCCGCACTCAGATCCTAGCGTCGAGCCCCCTTGAGTCAGGAAACATTTCAGACCTATGGA	70
p53.X54156	-----	70
p53.M14695	-----	70
p53.M14694	-----	70
p53.K03199	-----	70
 P53.WTPHP53C1	AAACTACTTCCGTAAAACAACGTTCTGTCCCCCTTGCCTCCAGAACATGGATGATTGATGCTGTCCCC	140
p53.X54156	-----	140
p53.M14695	-----	140
p53.M14694	-----	140
p53.K03199	-----	140
 P53.WTPHP53C1	GGACGATATTGAACAATGGTCACTGAAGACCCAGGTCCAGATGAAGCTCCCAGAACATGCCAGAGGCTGCT	210
p53.X54156	-----	210
p53.M14695	-----	210
p53.M14694	-----	210
p53.K03199	-----	210
 P53.WTPHP53C1	CCCCCCGTGGCCCTGCACCAGCAGCTCCTACACCGCGGCCCTGCACCAGCCCCCTTGGCCCTGT	280
p53.X54156	-----G-----	280
p53.M14695	-----GA-----	280
p53.M14694	-----G-----GA-----	280
p53.K03199	-----	280
 P53.WTPHP53C1	CATCTTCTGTCCCTTCCCAGAAAACCTACCAGGGCAGCTACGGTTCCGTCTGGCTTGCATTCTGG	350
p53.X54156	-----	350
p53.M14695	-----	350
p53.M14694	-----	350
p53.K03199	-----	350
 P53.WTPHP53C1	GACAGCCAAGTCTGTGACTTGCACGTACTCCCCTGCCCTCAACAAAGATGTTTGCCTGGCAAGACC	420
p53.X54156	-----	420
p53.M14695	-----	420
p53.M14694	-----	420
p53.K03199	-----	420

P53.WTPHP53C1	TGCCCTGTGCAGCTGTGGTTGATTCCACACCCCCGCCC GGCACCCCGTCCGCCATGGCATCTACA	490
p53.X54156	-----	490
p53.M14695	-----	490
p53.M14694	-----	490
p53.K03199	-----	490
P53.WTPHP53C1	AGCAGTCACAGCACATGACGGAGGTTGTAGGGCCTGCCCCCACC ATGAGCGCTGCTCAGATAGCGATGG	560
p53.X54156	-----	560
p53.M14695	-----	560
p53.M14694	-----	560
p53.K03199	-----	560
P53.WTPHP53C1	TCTGGCCCCTCCTCAGCATCTTATCCGAGTGGAAAGGAAATTGCGTGTGGAGTATTGGATGACAGAAC	630
p53.X54156	-----	630
p53.M14695	-----	630
p53.M14694	-----	630
p53.K03199	-----	630
P53.WTPHP53C1	ACTTTTCGACATAGTGTGGTGGTGCCTATGAGCCGCTGAGGTTGGCTCTGACTGTACCACCATCCACT	700
p53.X54156	-----	700
p53.M14695	-----	700
p53.M14694	-----	700
p53.K03199	-----	700
P53.WTPHP53C1	ACAACTACATGTGTAACAGTTCCCTGCATGGCGGCA TGAACCGGAGGCCATCCTCACCATCATCACACT	770
p53.X54156	-----	770
p53.M14695	-----	770
p53.M14694	-----	770
p53.K03199	-----	770
P53.WTPHP53C1	GGAAGACTCCAGTGGTAATCTACTGGGACGGAACAGCTTGAGGTGCGTGTGCGCTGCTGGAGA	840
p53.X54156	-----	840
p53.M14695	-----	840
p53.M14694	-----	840
p53.K03199	-----A-----	840
P53.WTPHP53C1	GACCGGCGCACAGAGGAAGAGAACTCCGCAAGAAAGGGGAGCCTCACCA CGAGCTGCCCGAGGAGCA	910
p53.X54156	-----	910
p53.M14695	-----	910
p53.M14694	-----	910
p53.K03199	-----	910
P53.WTPHP53C1	CTAACGCAGCACTGCCAACAACCCAGCTCCTCTCCCAGCCAAGAAGAAACC ACTGGATGGAGAATA	980
p53.X54156	-----	980
p53.M14695	-----	980
p53.M14694	-----	980
p53.K03199	-----	980
P53.WTPHP53C1	TTTCACCCCTCAGATCCGTGGCGTGAGCGCTTCGAGATGTTCCGAGAGCTGAATGAGGCCTTGGAACTC	1050
p53.X54156	-----	1050
p53.M14695	-----	1050
p53.M14694	-----	1050
p53.K03199	-----	1050
P53.WTPHP53C1	AAGGATGCCAGGCTGGGAAGGAGCCAGGGGGAGCAGGGCTCACTCCAGCCACCTGAAGTCCAAAAGG	1120
p53.X54156	-----	1120
p53.M14695	-----	1120
p53.M14694	-----	1120
p53.K03199	-----	1120
P53.WTPHP53C1	GTCAGTCTACCTCCGCCATAAAAACATGTTCAAGACAGAAGGGCTGACTCAGACTGA	1182
p53.X54156	-----	1182
p53.M14695	-----	1182
p53.M14694	-----	1182
p53.K03199	-----	1182

LOCUS Rb 4740 bp ss-mRNA PRI 15-JUN-1989
 DEFINITION Human retinoblastoma susceptibility mRNA, complete cds.
 ACCESSION M15400
 KEYWORDS retinoblastoma susceptibility.
 SOURCE Human fetal retina, cDNA to mRNA, clone RB-[1,5].
 REFERENCE 1 (bases 243 to 4740)
 AUTHORS Lee,W.H., Bookstein,R., Hong,F., Young,L.J., Shew,J.Y. and Lee,E.
 TITLE Human retinoblastoma susceptibility gene: Cloning, identification, and sequence
 JOURNAL Science 235, 1394-1399 (1987)
 REFERENCE 2 (bases 1 to 480)
 AUTHORS Lee,E., Bookstein,R., Young,L.J., Lin,C.-J., Rosenfeld,M.G. and Lee,W.H.
 TITLE Molecular mechanism of retinoblastoma gene inactivation in retinoblastoma cell line Y79
 JOURNAL Proc. Natl. Acad. Sci. U.S.A. 85, 6017-6021 (1988)
 COMMENT Draft entry and computer-readable copy of sequence in [1] kindly provided by R.Bookstein, 27-APR-1987.

The authors [1] identified the retinoblastoma susceptibility (RB) gene encoding a messenger RNA (mRNA) of 4.6 kb on the basis of chromosomal location, homozygous deletion, and tumor-specific alterations in expression. Transcription of the gene was abnormal in 6/6 retinoblastomas while normal in fetal retina and placenta. The sequence presented in this entry is derived from human fetal retina cDNA. The authors [2] also examined cDNA from the retinoblastoma cell line Y79. They found that the mRNA produced was a shortened transcript which reflects a deletion of exons 2-6 and must have occurred by a mechanism other than recombination of homologous sequences.

The Retinoblastoma (Rb) susceptibility gene maps to chromosome 13q14 and encodes a nuclear phosphoprotein of 105kDa. Rb is a tumor suppressor gene; its nuclear location and ability to bind DNA suggests a role in transcriptional regulation. On the side of the protein lies a complementary oncoprotein-binding pocket that is a common target for the three viral proteins: E1A of adenovirus, large T antigen of SV40, and E7 of HPV. Munger et al. (EMBO 8, 4099-4105) determined that the high risk HPV types such as HPV16 and HPV18 bind to pRB with higher affinity than low risk types 6b and 11. The amino acid locations of the pRB binding site of the HPV16 E7 protein have been mapped to a small stretch of amino acids. In addition, pRb has shown to associate with a number of cellular host proteins at the onco-protein binding site. pRB shows differential phosphorylation at different points within the cell cycle. All three of the viral proteins mentioned above interact with the underphosphorylated form of pRB.

BASE COUNT 1508 a 887 c 862 g 1483 t
 ORIGIN 1038 bp upstream of EcoRI site; chromosome 13q14.1-q14.2.
 1 ttccggtttt tctcaggggaa cgttgaatt attttgtaa cgggagtcgg gagaggacgg
 61 ggcgtgcccc gcgtgcgcgc gcgtcgctt ccccggegt cctccacagc tcgctggctc
 121 ccggccggaa aaggcgcat gccgccccaa acccccccggaa aaacggccgc caccggccgc
 181 gctgccggcg cgaaaccccc ggacccggcg ccggccggcc ctcttgagga ggacccagag
 241 caggacagcg gcccggagga cctgccttc gtcaggctt agtttgaaga aacagaagaa
 301 cctgattta ctgcattatg tcagaaatta aagataccag atcatgtcag agagagagct
 361 tggtaactt gggagaaagt ttcatctgtg gatggagttat tattcaaaag
 421 aaaaaggaac tggggaaat ctgtatctt attgcacggat ttgacccatgat tgatgtcg
 481 ttcactttac tgagctacag aaaaacatac gaaatcgtg tccataaaatt cttaactta
 541 ctaaaaagaaa ttgataccag taccaaaggataatgatgtcaagact gttgaagaag
 601 tatgtatgtat tggatgcact cttcagcaaa ttggaaaggat catgtaaact tatataatttgc
 661 acacaaccca gcagttcgat atctactgaa ataaattctg cattgggtct aaaagttct
 721 tggatcacat tttttagc taaaggggaa gtattacaatgatga tctgggtatt
 781 tcatttcagt taatgtatg tgccttgac tattttatata aactctcacc tcccatgttg

Retinoblastoma

841 ctc当地aaac catataaaac agctgttata cccattaatg gttcacctcg aacaccagg
901 cgaggcaga acaggagtgc acggatagca aaacaactag aaaatgatac aagaattatt
961 gaagttctct gt当地aaaca tgaatgtat atagatgagg tgaaaaatgt ttattncaaa
1021 aattttatac ctttatgaa ttctcttggc ct当地taacat ctaatggact tccagagggt
1081 gaaaatctt ctaaacgata cgaagaaatt tatcttaaaa ataaagatct agatcgaaga
1141 ttattnnccg atcatgataa aactcttcg actgattcta tagacagttt tgaaacacag
1201 agaacaccac gaaaagttt ctttgatgaa gaggtgataa taattccctcc acacactcca
1261 gtttagactg tt当地aacac tatccaacaa ttaatgtatgaa ttttaattt tgc当地gtat
1321 caacccctcg aaaatctgtat ttcccttattt aacaactgca cagtaatcc aaaagaaagt
1381 atactgaaaa gagtgaagga tataggatac atctttaaag agaaatttgc taaagctgt
1441 ggacagggtt gt当地gaaat tggatcacag cgatacaac ttggagttcg ct当地tattac
1501 cgagtaatgg aatccatgtat taaatcgaa gaagaacat tatccattca aaatttttagc
1561 aaacttctgtat atgacaacat tttcatatgat ttttatttgg cgtgc当地tct tgagggtgta
1621 atggccacat atagcagaag tacatctcg aatcttgatt ct当地acaga ttttgc当地
1681 ccatggattc tgaatgtgc taattttaaa gc当地tttatttgc tttacaaaagt gatc当地aaagt
1741 tttatcaaag cagaaggca cttgacaaga gaaatgataa aacatttgc acgatgtgaa
1801 catgaatca tggatccct tgc当地ggctc tc当地atttgc ct当地tatttgc tcttattt
1861 caatcaaagg accgagaagg accaactgat caccttgc当地 ct当地gttcc tcttattt
1921 cctctccaga ataatcacac tgc当地ggatgat atgtatctt ct当地gttcc agtccaaag
1981 aaaaaaggtt caactacgca cttgaaatttct actgcaatgc cagagacaca agc当地acctca
2041 gc当地ttccaga cccagaagcc attgaaatct acctcttcc ct当地gttcc taaaaaaatgt
2101 tattccggatc cctatctccg gctaaataca ct当地gttcc gcttccgtc tgacccacca
2161 gaatttgc当地 atatcatctg gacccttttgc cagcacaccc tgc当地gttcc gatgactc
2221 atgagagaca ggc当地tttgc当地 ccaatttgc当地 atgtgttcc tgc当地gttcc atgcaatgt
2281 aagaatatag accttatttgc当地 ctttgc当地tgc当地tgc当地tgc当地tgc当地
2341 gttcaggaga cattcaaagc ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
2401 tt当地tataact cggcttccat gcaaggactg aaaacaaaata ttttgc当地tgc当地tgc当地
2461 aggcccccttccat ctttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
2521 tc当地cccttccat gggatccctgg agggacatc tatatttgc当地tgc当地tgc当地tgc当地
2581 atttgc当地atgatc accaacaatggc atgacttccat gatcaagaat ct当地gttcc
2641 atttgc当地atgatc cattcgggac ttcttgc当地tgc当地tgc当地tgc当地tgc当地
2701 agc当地accgtg ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
2761 ctacgttccat atatttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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2881 aaaaatgtatgatc taccttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
2941 ggtggacttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3001 cccagggttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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3121 acaaggatc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3181 cttccaaatgtatgatc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3241 tgatc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3301 ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3361 ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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3601 acagatccat taccttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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3781 aggcccttccat ctttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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3901 tt当地tatttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
3961 gtttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4021 ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
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4141 tatcttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4201 tccccc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4261 atgcttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4321 ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4381 ttttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4441 ctttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4501 agtttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4561 tagtgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地
4621 agatatttgc当地tgc当地tgc当地tgc当地tgc当地tgc当地tgc当地

Retinoblastoma

4681 gttactattt tctacaatta atagtttgc tatTTTaaa aaaaTTTTtt gttaaggatc

Deleted in Colon Cancer

LOCUS DCC 4608 bp ss-mRNA PRI 04-FEB-1994
DEFINITION Homo sapiens DCC mRNA.
ACCESSION X76132
KEYWORDS cell adhesion molecule; DCC gene; immunoglobulin gene superfamily; transmembrane protein; tumor suppressor gene.
SOURCE Human normal adult and fetal brain, Clontech cDNA libraries
CAT #HL1065b, HL1003b.
REFERENCE 1 (bases 1 to 4608)
AUTHORS Cho, K.
TITLE ;
JOURNAL direct submission
REFERENCE 2 (bases 1 to 4608)
AUTHORS Hedrick L., Cho K.R., Fearon E.R., Wu T.C., Kinzler K.W., Vogelstein B.
TITLE The DCC gene product in cellular differentiation and colorectal tumorigenesis
JOURNAL Unpublished
COMMENT Tumors were induced in nude mice after transplantation of the HPV-18 immortalized human keratinocyte cell line (1811) and treatment with the carcinogen nitrosomethylurea (NMU). In these transformed cells, one allele of the deleted in colon cancer DCC tumor-suppressor gene was absent and the other allele was mutated (Klingelhutz et al. Oncogene 8: 95-9).
Related sequence: M32292
BASE COUNT 1282 a 1179 c 1058 g 1089 t
ORIGIN
1 atggagaata gtcttagatg tgtttggta cccaagctgg cttttgtact ctccggagct
61 tccttgctca gcgcgcacatc tcaagtaacc ggaaaaaaa ttaaagcttt cacagcactg
121 cgcttcctct cagaaccttc tgatgccgtc acaatgcggg gagaaatgt cctcctcgac
181 tgctccgccc agtccgaccg aggagttcca gtatcaagt ggaagaaaga tggcattcat
241 ctggccttgg gaatggatga aaggaagcac caactttca aatgggtctct gctgatacaa
301 aacataacttc attccagaca ccacaaggca gatgagggac ttaccaatg tgaggcatct
361 tttaggagat ctggctcaat tattatgcgg acagcaaaag ttgcgttagc aggaccactg
421 aggttccctt cacagacaga atctgtcaca gccttcatgg gagacacagt gctactcaag
481 tgtgaagtca ttggggagcc catgccaaca atccacttggc aagaagacca acaagacctg
541 actccatcc caggtgactc ccggatgggt gtcttgcctt ctggagcatt gcagatcagc
601 cgactccaac cgggggacat tggaaattac cgatgctcag ctcgaaatcc agccagctca
661 agaacaggaa atgaagcaga agtcagaatt ttatcagatc caggactgca tagacagctg
721 tattttctgc aaagaccatc caatgttagta gccattgaag gaaaagatgc tgcctggaa
781 tgggtgttt ctggctatcc tccaccaagt ttacctgtt tacgaggcga ggaagtcac
841 caactcaggat ctaaaaagta ttctttattt ggtgaaagca acttgcatt ctccaatgt
901 acagatgtat acagtggaaat gtatacctgt gttgtcacat ataaaaatga gaatattatgt
961 gcctctgcag agtcacagt ctgggttccg ccatgggtt taaatcatcc ttccaacctg
1021 tatgcctatg aaagcatgaa tattgagttt gaatgtacag tctctggaaa gcctgtgccc
1081 actgtgaatt ggatgaagaa tggagatgt gtcattccta gtgattatcc tcaaatgt
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4561 aacaaaagca aagatgcatt ttcactgaa tgtcaaagtt tagctgtct

E2F1

LOCUS E2F1 2517 bp ss-mRNA PRI 10-AUG-1992
DEFINITION Homo sapiens (E2F-1) pRB-binding protein mRNA, complete cds.
ACCESSION M96577
KEYWORDS DNA-binding protein; pRB-binding protein; transcription factor E2F.
SOURCE Homo sapiens fetal brain cDNA to mRNA.
REFERENCE 1 (bases 1 to 2517)
AUTHORS Helin,K., Lees,J.A., Vidal,M., Dyson,N.J., Harlow,E. and Fattaey,A.
TITLE A cDNA encoding a pRB-binding protein with properties of the transcription factor E2F
JOURNAL Cell 70, 337-350 (1992)
COMMENT E2F, a cellular transcription factor, forms an inactive complex with pRB (retinoblastoma tumor suppressor protein). HPV-16 E7 is one of several DNA virus oncoproteins that dissociates the E2F-pRB complex. This dissociation is coupled with enhanced E2F-dependent transcription. In the S-phase of the cell cycle, E2F complexes with cyclin A, p107, and cdk2 kinase. Arroyo et al. (Mol and Cell Bio 13: 6537-46) report that the E7 protein of HPV-16 associates with the E2F-cyclin A complex. Arroyo et al. (Mol and Cell Bio 13: 6537-46) further note that a high risk HPV has a higher binding efficiency to the E2F-cyclin A complex than the low-risk HPV type. Pagano et al. (Oncogene 7: 1681-6) report that the E2F-cyclin A complex can occur in HPV-18 infected cells independent of pRB.
BASE COUNT 454 a 784 c 781 g 498 t
ORIGIN
1 ggaattccgt ggccggact ttgcaggcag cggcgccgg gggcgagcg ggatcgagcc
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E2F1

2401 cccactgctc tgccccaccc tccaatctgc actttgattt gcttcctaac agctctgttc
2461 cctcctgctt tggtttaat aaatatttg atgacgttaa aaaaaggaat tcgatat

Epidermal Growth Factor Receptor

LOCUS EGFR 5532 bp ss-mRNA PRI 13-JUN-1985
DEFINITION Human mRNA for precursor of epidermal growth factor receptor
ACCESSION X00588
KEYWORDS epidermal growth factor receptor; signal peptide.
SOURCE
REFERENCE 1 (bases 1 to 5532)
AUTHORS Ullrich A., Coussens L., Hayflick J.S., Dull T.J., Gray A.,
Tam A.W., Lee J., Yarden Y., Libermann T.A., Schlessinger J.,
Downward J., Mayes E.L., Whittle N., Waterfield M.D., Seuberg P.H.;
TITLE Human epidermal growth factor receptor cDNA sequence and aberrant
expression of the amplified gene in A431 epidermoid carcinoma
cells
JOURNAL Nature 309:418-425(1984).
COMMENT Cohen et al. demonstrated that the BPV E5 protein activates the
EGF receptor through complex formation. They have further
determined that this activation is specific to the cytoplasmic
domain of the EGF receptor and is ligand independent (J Virol
67: 5303-11). Straight et al. (J Virol 67: 4521-32) demonstrated
that E5 HPV-16 infected keratinocytes had a two to five fold
increase in EGF receptors.

BASE COUNT 1472 a 1484 c 1337 g 1239 t
ORIGIN

1 gccgcgtgc gcccggatcc cgagcttagcc ccggcgccgc cgccgccccag accggacgac
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5401 cacacacaca tacaatgtt tccttttgc tttaaatgtt ttttgcactc ccagatc
5461 cagggccctt acagcattgtt taagaaaatgtt tttgatgttgc tctcaatgc aaataaaact
5521 atattcattt cc

ERBB2

LOCUS ERBB2 4473 bp ss-mRNA PRI 30-JUN-1987
DEFINITION Human c-erb-B-2 mRNA.
ACCESSION X03363
KEYWORDS c-myc proto-oncogene; cell surface glycoprotein; erbB gene; erbB oncogene; glycoprotein; growth factor; growth factor receptor; kinase; neu oncogene; transmembrane protein; tyrosine kinase.
SOURCE Homo sapiens cell line MKN-7
REFERENCE 1 (bases 1 to 4473)
AUTHORS Yamamoto,T., Ikawa,S., Akiyama,T., Semba,K., Nomura,N., Miyajima,N., Saito,T. and Toyoshima,K.
TITLE Similarity of protein encoded by the human c-erb-B-2 gene to epidermal growth factor receptor
JOURNAL Nature 319, 230-234 (1986)
COMMENT Milde-Langosch et al. (Verh Dtsch Ges Pathol 75: 363-5) demonstrated an increase in expression of c-erbB2 in 4/11 HPV-positive endometrial carcinomas and in 3/19 HPV-positive CIN3 lesions.

The c-erb-B-2 protein shows similarity to the epidermal growth factor receptor.

EMBL features not translated to GenBank features:

key	from	to	description		
SITE	376	384	pot. glycosylation site		
SITE	544	558	pot. glycosylation site		
SITE	733	741	pot. glycosylation site		
SITE	949	957	pot. glycosylation site		
SITE	1762	1770	pot. glycosylation site		
SITE	1885	1893	pot. glycosylation site		
SITE	2059	2067	pot. glycosylation site		
SITE	2353	3132	aa 727-986, seq. homologous to EGF receptor kinase domain		
SITE	2446	2454	pot. glycosylation site		
SITE	4455	4460	put. polyA signal		
POLYA	4473	4473	polyA site		
BASE COUNT	902	a 1383	c 1329	g 859	t

ORIGIN

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K-ras

LOCUS KRAS 5775 bp ds-DNA PRI 05-JUN-1991
DEFINITION Human K-ras oncogene, complete cds.
ACCESSION M54968 M38506
KEYWORDS K-ras oncogene.
SOURCE Human tumor DNA.
REFERENCE 1 (bases 1 to 5775)
AUTHORS Kahn,S., Yamamoto,F.-I., Almoguera,C., Winter,E., Forrester,K., Jordano,J. and Perucho,M.
TITLE The c-K-ras gene and human cancer (review)
JOURNAL Anticancer Res. 7, 639-652 (1987)
BASE COUNT 1739 a 974 c 1105 g 1957 t
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K-ras

5761 taaatcatta ccagg

**IV-18
SEP 94**

Monocyte Chemoattractant Protein

LOCUS MCP1 3227 bp ds-DNA PRI 15-JUN-1994
DEFINITION Human MCP-1 gene for monocyte chemoattractant protein-1.
ACCESSION D26087
KEYWORDS monocyte chemoattractant protein-1.
SOURCE Homo sapiens (library: EMBL3 SP6/T7) placenta DNA.
REFERENCE 1 (sites)
AUTHORS Rollins,B.J., Stier,P., Ernst,T. and Wong,G.G.
TITLE The human homolog of the JE gene encodes a monocyte secretory protein
JOURNAL Mol. Cell. Biol. 9, 4687-4695 (1989)
REFERENCE 2 (sites)
AUTHORS Shyy,Y.-J., Li,Y.-S. and Kolattukudy,P.E.
TITLE Structure of human monocyte chemotactic protein gene and its regulation by TPA
JOURNAL Biochem. Biophys. Res. Commun. 169, 346-351 (1990)
REFERENCE 3 (bases 1 to 3227)
AUTHORS Ueda,A., Okuda,K., Ohno,S., Shirai,A., Igarashi,T., Matsunaga,K., Fukushima,J., Kawamoto,S., Ishigatubo,Y. and Okubo,T.
TITLE NF-kB and Sp1 regulate transcription of human monocyte chemoattractant protein-1 gene
JOURNAL J. Immunol. (1993) In press
COMMENT Rosl et al. (J Virol 68: 2142-50) demonstrated that MCP-1 protein is not expressed in HPV-18 positive cervical carcinoma cells, despite the fact that the gene which encodes it, the JE gene, has no mutations. Conversely, in non-malignant cells the tumor necrosis factor alpha induces MCP-1 expression which results in HPV-18 repression.

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NCBI gi: 516772
BASE COUNT 850 a 793 c 732 g 852 t
ORIGIN

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Monocyte Chemoattractant Protein

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2701 ttgcgttcac agaaagcaga atcctaaaaa ataaccctct tagttcacat ctgtggcag
2761 tctgggctta atggcaccccc atcctccca ttgcatttgc tggcgtcagc agtgaatgg
2821 aaaagtgtct ctgcatttgc accctgcttcc ctgcattactt tcctgaaat ccacaggatg
2881 ctgcatttgc tcagcagatt taacagccca ctatcactc atggaaagatc cctccctcctg
2941 ctgcactccg ccctctctcc ctgcggccgc ttcaataag aggcagagac agcagccaga
3001 ggaaccgaga ggctgagat aaccctagaaa catccaattc tcaaactgaa gctcgactc
3061 tcgcctccag catgaaagtc tctgcggcc ttgcatttgc gctgcata gcagccacat
3121 tcattccca agggctcgct cagccaggtt agggccccctt ttgcatttgc tgaaccacat
3181 tgtcttcctt ctgagttatc atggaccatc caagcagacg tggtaacc

LOCUS NF1 3238 bp ss-mRNA PRI 02-NOV-1990
 DEFINITION Human hepatic nuclear factor 1 (HNF1) mRNA, complete cds, clones HCL10, HCL12, HCL17, and HCL20.
 ACCESSION M57732 J04771
 KEYWORDS hepatic nuclear factor 1; transcription factor.
 SOURCE Homo sapiens liver NF1 cDNA to mRNA.
 REFERENCE 1 (bases 1 to 3238)
 AUTHORS Bach,I., Galcheva-Gargova,Z.I., Mattei,M.-G., Simon-Chazottes,D., Guenet,J.-L., Cereghini,S. and Yaniv,M.
 TITLE Cloning of human hepatic nuclear factor 1 (HNF1) and chromosomal localization of its gene in man and mouse
 JOURNAL Genomics 8, 155-164 (1990)
 COMMENT Chong et al. (Nucleic Acids Res 18: 465-70) report that the HPV-16 enhancer is activated by NF1, AP-1 and TEF-2, and that it occurs without cooperation. Taniguchi et al. (Virology 195: 500-10) demonstrated reduced enhancer activity upon mutation of any of the three NF-1 binding sites (GCCAA).

BASE COUNT 651 a 1100 c 955 g 531 t 1 others
 ORIGIN

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1 cgtggccctg tggcagccga gccatggttt ctaaacttag gccaatgcac acggagactcc
61 tggcgccctt gctcgagtca gggctgagca aagaggcact gatccaggca ctgggtgagc
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661 tcctgttcca ggcctatgg aggcagaaga accctagaa ggaggagcga gagacgctag
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Nuclear Factor 1

2701 cggggctggg aagtgcgtcct tactcctgtg ggagcctcg aaccctgtcc aagtccaggt
2761 cctgggtgggg cagtcctct gtctcgagcg ccctgcagac cctgccttg tttggggcag
2821 gagtagctga gtcacaagg cagcaaggcc cgagcagctg agcagggccg gggactggc
2881 caagctgagg tgcccaggag aagaaagagg tgaccccagg gcacaggagc taactgtgt
2941 gacaggacta acactcagaa gcctgggtgc ctggctggct gaggcagtt cgcagccacc
3001 ctgaggagtc tgaggtcctg agcaactgcca ggagggacaa aggacctgt gaacctcagga
3061 caagcatggt ccacatccc tggcctgtct gctgagaacc tggccttcag tgtaccgcgt
3121 ctaccctggg attcaggaaa aggcctgggg tgacccggca cccccctgcag cttgttagcca
3181 gcccgggcga gtggcacgtt tatthaactt ttagtaaagt caaggagaaa tgccgtgg

LOCUS OCT1 3824 bp ss-mRNA PRI 18-NOV-1993
 DEFINITION Human octamer binding transcription factor 1 (OTF1) mRNA, complete
 cds.
 ACCESSION L20433
 KEYWORDS octamer binding transcription factor 1.
 SOURCE Homo sapiens (library: lambda pSH4K) cDNA to mRNA.
 REFERENCE 1 (bases 1 to 3824)
 AUTHORS Bhargava,A.K., Zhen,L.I. and Weissman,S.M.
 TITLE Differential expression of four members of the POU family of
 proteins in activated and TPA-treated Jurkat T-cells
 JOURNAL Proc. Natl. Acad. Sci. U.S.A. 93, 10260-10264 (1994)
 COMMENT Hoppe-Seyler et al. (J Virol 65: 5613-8) demonstrated that Oct-1
 can repress transcription of HPV-18. Mutagenesis experiments
 suggest that this effect is mediated without direct DNA binding.
 Morris et al. determined that HPV-16 transcription is repressed
 by Oct-1 only in non-cervical cells. Conversely, in cervical
 cells Oct-2 is expressed which transactivates HPV-16 transcription.
 BASE COUNT 896 a 1005 c 1067 g 856 t
 ORIGIN
 1 gcggggctag agctgtcgg a aagcgccgac cgcgaggccg ggcgcggccg ctctgcgg
 61 tcagagggag cgccctggcag c a caggagc a g c a g c a g c a g c a g c a g c
 121 c a g c c c c g c g a c c g c c g c g g t c a g c a c t c c g a a g g g
 181 c a c t t c c c g c g g a c t t c g g a g t g t t g g a g t a t a c a t
 241 t c c a t g a a c a g c a g c c t c a c t t t g c a t g c a t c c c a
 301 c c g t c g c t c g a c t c c a g c t c c g a g g c c a t c g c t g c
 361 c a g a g a a c c t c t c g c c a g c t g g a c g a g a c g c t g
 421 g c c g t g g a c a t c g c c a g g g c a a g a g c a t c t t t c
 481 c a c a c g a t g a a c a g c g t g c a c g t c c a t t c c a c
 541 c a c c a c c a c c a c c a c c a g c t c g a a c c c g c t g
 601 c c g t c g c t c g c g c t c a t g g c c g g g c g g c g c t g
 661 g c c c a c g a c g g c c g g g g g g g g g g g g g g g g
 721 g g c c c c g
 781 c t c c t g g c g g c a c c c c c a c c c a c a t g c a
 841 c c c g c g g c g g c a c a c a c a c a c a c a c a c
 901 g c g g c g c g c g c a c a c a c a c a c a c a c a c
 961 g c g g c a t c g g c g g c g g c g g c g g c g g c
 1021 g a c a c g g a c c g c g a g a c t c g g a g c
 1081 c t g g g c g t g a c g a g g c g a c t g g g c t c
 1141 g g c t c a c t c a c a g a g c a c a c a c a c a c
 1201 a t g a t c g c g t c a a g c c c a t c g a g g c
 1261 g a g a a a t g a a c a g c c t g a c t t c a a g
 1321 a t c g c c g c g c g a a g c g c g a g g c
 1381 t c c g a g a a g a t c g c c g c c a g a a a
 1441 t g g t t t g c a a c c a g a g a a g c
 1501 g g c t g g g a g g t c g g c g g a c a g a a t
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 1621 c c t t c t c g c t c t c c a c c c a t t c
 1681 c t t g a g g a c t g g g a g g a g g a g g
 1741 g c a t t c g g g g a g g t g g g a g g a g g
 1801 g a g c a g g a t g t t c t g g g g g g
 1861 g t t t c t g a c c a g a c a c t t
 1921 a a a c c t a c c a a a a c t a g a c
 1981 a g c a a a g g a a a t g c t t a g
 2041 t c c c c c a t a g a a g a g a a a a
 2101 a c g a a c t g a g c a a a a a c
 2161 g c t a g t g t g t g t g t a t t
 2221 c t c t t a t t c c t c g t t c
 2281 g t g g g a g c a c a g g a a a g g
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 2461 a t c t t a t t c a g a a a a t g a
 2521 a t c t a g t t t a g a c a t t c
 2581 t t c a g t t t a t t a a a a t
 2641 a t a g a c t a a t a g g a g

Oct1

2701 gagactcatc ttgtcttct aggtcccggtt tcttcctctc ttggaggaca tgaaattata
2761 gaaatgttga gaagttcctg ctttcttttg cggttaggact tggctgtgag aaaatcacct
2821 aaatcccaga aaagaggaag acagattaa agtgc(ccca cccccatttg ttcaaagag
2881 gtctgcatgt tggcgaaaa cagaacaact gtgttcctt ttacttgttca ttattattca
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Platelet Derived Growth Factor Receptor

LOCUS PDGFR 5570 bp ss-mRNA **PRI** 28-SEP-1992
DEFINITION Human platelet-derived growth factor (PDGF) receptor mRNA, complete cds.
ACCESSION J03278
KEYWORDS cell surface glycoprotein; glycoprotein; kinase; tyrosine kinase.
SOURCE Human skin fibroblast cell (SK5), cDNA to mRNA, clone pHPDGFR.
REFERENCE 1 (bases 1 to 5570)
AUTHORS Gronwald,R.G.K., Grant,F.J., Haldeman,B.A., Hart,C.E., O'Hara,P.J., Hagen,F.S., Ross,R., Bowen-Pope,D.F. and Murray,M.J.
TITLE Cloning and expression of a cDNA coding for the human platelet-derived growth factor receptor: evidence for more than one receptor class
JOURNAL Proc. Natl. Acad. Sci. U.S.A. 85, 3435-3439 (1988)
COMMENT Nilson and DiMaio (Mol Cell Biol 13: 4137-4145, Cohen et al. J Virol 67:5303-11) reported that the BPV-1 E5 protein activates PDGF beta receptors through complex formation resulting in fibroblast transformation. Cohen et al. demonstrated that this activation is specific to the transmembrane domain of the PDGF receptor and that it is ligand independent (Cohen et al. J Virol 67: 5303-11).
 Draft entry and computer-readable sequence [1] kindly submitted by R.G.K.Gronwald, 06-APR-1988.
BASE COUNT 1195 a 1676 c 1530 g 1169 t
ORIGIN Chromosome 5q31-q32.

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Platelet Derived Growth Factor Receptor

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2521 ttgggctccc cctgcccagc catgtgcct tgaccgggaa gagcgcacggt ggctacatgg
2581 acatgagcaa ggacgagtcg gtggactatg tgcccatgct ggacatgaaa ggagacgtca
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4561 ttggacactgc tatgaggctt tggaggaatc ctcacccttc tctggccctc agttccct
4621 tcaaaaaatg aataagtctgg attttaac tctgagtgcc ttgccagcac taacattcta
4681 gatgttcca ggtgggtgc catttgtcca gatgaagca ggcataatac cctaaacttc
4741 catctgggg gtcagctggg ctccctggag attccagatc acacatcaca ctctggggac
4801 tcaggaacca tggcccttcc ccaggccccca agcaagtc tcaaaacacag ctgcacaggc
4861 ctgcacttag agtgcacagcc ggtgtcctgg aaagccca gcaagtcctc cagggacatg
4921 ggaagaccac gggacacttcc tcaactacca cgtacacttc cgggggttac ctggccaaa
4981 gggacaaga gggcaaatga gtcacaccc tgcagccac cactccagca cctgtgcga
5041 ggtctgcgtc gaagacagaa tggacagtga ggacagtat gttttgtaaa agacaagaag
5101 cttcagatgg taccccaaga aggtatgttag aggtggccgc ttggagttt ccctcaccc
5161 accagctgc ccattccctga ggcagcgtc catggggta tgggtttgtc actgcccaga
5221 ctcaggatgtt acatctcatt tggccctggcc cagtggccat tggaggtgccc agggggatca
5281 ggggtttagc caagacgccc cccgcacgggg agggttggga aggggggtgca ggaagctcaa
5341 cccctctggg caccacccctt gcaattgcagg ttggcacctt acttccctgg gatccccaga
5401 gttgtccaa ggagggagag tgggttctca atacggtacc aaagatataa tcaccttagt
5461 ttacaaatat ttttaggact cactttaact cacattata cagcagaaat gctatggat
5521 atgctgtttaa gttttctat ctgtgtactt ttttttaagg gaaagat

PP2A 72kDa Regulatory Subunit

LOCUS PP2A72 2338 bp ss-mRNA PRI 20-AUG-1993
DEFINITION Homo sapiens protein phosphatase 2A 72 kDa regulatory subunit mRNA, complete cds.
ACCESSION L12146
KEYWORDS phosphoprotein phosphatase 2A; phosphoprotein phosphohydrolase; protein phosphatase 2A 72 kDa regulatory subunit; regulatory subunit.
SOURCE Homo sapiens (library: lambda ZAP; Stratagene) female heart muscle cDNA to mRNA.
REFERENCE 1 (bases 1 to 2338)
AUTHORS Hendrix,P., Mayer-Jaekel,R.E., Cron,P., Goris,J., Hofsteenge,J., Merlevede,W. and Hemmings,B.A.
TITLE Structure and expression of a 72-kDa regulatory subunit of protein phosphatase 2A. Evidence for different size forms produced by alternative splicing
JOURNAL J. Biol. Chem. 268, 15267-15276 (1993)
COMMENT PP2A consists of regulatory subunits and one catalytic subunit. In their review, "Protein Phosphatase 2A and the Regulation of Human Papillomavirus Gene Activity, Schegget and Noordaa (in Human Papillomaviruses, 121-9) report that PP2A is inactivated by high expression of its regulatory subunit, PR55Beta. This inhibition is linked to the activation of the LCR region of HPV-16. Thus, the authors suggest that the regulation of the HPV-16 promoter is correlated to cellular protein phosphorylation.

BASE COUNT 694 a 497 c 526 g 621 t
ORIGIN

1 gaattccgca gctgtgttta taacttgaaa attatgagca ggtattggga gccacagact
61 gtatttggga acagctgcat gtcaggacaa gcatacgattt tggtctgtgg aggctgggag
121 agcacaatgc ttgaattca tttctttggg agataagaag aaaaaatacc ttcttacctg
181 acaagatgtca caacttgcta aatctgttata tcttcagaag tggctggac agttgcgaag
241 taaacattta ctttcaaaag atgatgttca aggaaacatc tctacgaaagg gacccggatt
301 taaggggaga gctagcttc ctggcaaggg gctgttgattt tggttccct tcacggttta
361 agaaggcggt gaagtctttt cagcagcacac agattcaaaa taaaccagaa aagaaacctg
421 gaacaccaact cccaccttca gcccaccttc caagtagtcc ccgaccccttc tccccggttc
481 cccatgtgaa taatgttgc aatgcgcatt tggccataaaa cattccacgg ttctactttc
541 ctgaaggact cccagatacc tggatgttcaatc atgaacaaatc tctaagcaga attgaaaactg
601 ctttcatggaa tattgaagaa cagaaaggcag acattttatga aatggggaaa attgcaaagg
661 tctgtggctg tcctcttat tggaaagcc ccatgttcag ggctgcagggg ggagagaaga
721 caggattttt gacagcacag tcattcattt ccatgtgggg aaagttgtg aataaccatc
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841 aggaggattt catccctcta cttcaggatg tggtgatc ccacccgtt ctcacgttcc
901 tgaaagatgc tccagaattt cactcccgct acatcaccac ggttatttc agaatattt
961 acacagtcaa cagatcttgg agtggaaaaa ttacttcgac agagataaga aaaagcaact
1021 ttttgcacaaac cctagcactt ttggagaagag aggaagatataa aacccaaattt acagattact
1081 tctccatgtt acatttctat gtttattttt gtaaattctg ggaacttagat actgatcacg
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1261 gaatgagctt tgcagatttt gtttgggtttt tgatctctg agaagacaaa aggaatccctt
1321 ccagcattgtt gtattgggtt cgcgtcatgg atgtggatgg agacgggttta ctctccatgt
1381 atgagcttggaa gtacttctat gaggagcagt gtgaacggat ggaaggccatg ggaatttggc
1441 ccttgcattt ccatgttta ctgtgccaga tgcttgacat agtgaagcca gctgttgatg
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1561 tctttatctt ggagaaataac tttagaccatg aacagagaga tccctttgcg gtcagaagg
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1681 agacgcttgtt tgcagggaa tctggccaaag cacaatttca ggaagggtttt gaagattatg
1741 aaacagatgtt acctgcctt ccctctgtt atttttttttt aagcaataaaa atattaatgt
1801 caagcttcc agagaaatgtt gggaaatgtt aatcagtgtt gtaagaatag ctggccgtt
1861 ctacaatgtt acgaagatgtt gtatctttttt ttgttgcgtt atgttctcgat
1921 ttgcatactt cttttttttt aatctttttt ctccaaatgtt gtatcatctg cactaggaaac
1981 ttttttttttta agcaataggtt ctggatatacc atttttttttt ttgttgcgtt
2041 tcaaaacctt tacggatgtt ctcctcgtt gttttttttt ttgttgcgtt
2101 ctacacttccatgtt gttttttttt gttttttttt gttttttttt
2161 aggtgtactgtt gttttttttt gttttttttt gttttttttt gttttttttt

PP2A 72kDa Regulatory Subunit

2221 agcctagaag aaaccccac ttgcagaaaa cttgagtcag aaaattctgg aacttgaaaa
2281 agtagtaagg gcctccagaa ttgacttagc cctagtaata aaagcaactgc cggaattc

PP2A Catalytic Subunit

LOCUS PP2ACAT 2966 bp ds-DNA PRI 04-MAR-1991
DEFINITION Human protein phosphatase 2A catalytic subunit-alpha gene, complete cds.
ACCESSION M60483 J05297
KEYWORDS protein phosphatase-2A catalytic subunit-alpha.
SOURCE Human placenta leukocyte DNA.
REFERENCE 1 (bases 1 to 2966)
AUTHORS Khew-Goodall,Y., Mayer,R.E., Maurer,F., Stone,S.R. and Hemmings,B.A.
TITLE Structure and transcriptional regulation of protein phosphatase 2A catalytic subunit genes
JOURNAL Biochemistry 30, 89-97 (1991)
COMMENT PP2A consists of regulatory subunits and one catalytic subunit. In their review, "Protein Phosphatase 2A and the Regulation of Human Papillomavirus Gene Activity, Schegget and Noordaa (in Human Papillomaviruses, 121-9) report that PP2A is inactivated by high expression of its regulatory subunit, PR55Beta. This inhibition is linked to the activation of the LCR region of HPV-16. Thus, the authors suggest that the regulation of the HPV-16 promoter is correlated to cellular protein phosphorylation.
Although this is a genomic sequence, the introns have been omitted. Intronic sequences have been requested. The data is presented as submitted by the author.
NCBI gi: 190223
BASE COUNT 747 a 728 c 732 g 759 t
ORIGIN
1 aaccacccggc gaggagcggg gcgcgtggaa gcgagcccg gtccgaggcc caaagaaaag
61 cccaaagcctc gccccccgcca tcgcgcccga cgagacacct aggtccgggg acgggtgtgt
121 gccgcggaaag tcaggtgcac tgcgcagcac tcccccggtt ggtacacgct cctccaccta
181 cgagtgcacct aattacaagg tgccagccgc gcccagaggt ggggggtggtt aatccaaagcg
241 gccactcgct gcccgttcct gcccccaaaag atgacggaaa cccacacat tacagagccg
301 cagcacccca gatgagccac ggggtcgcaa ttctcgttt cgtgatcgga ctgccaggcc
361 ccaggtgagg agctgagttc atcaggacag cggccitccc agggaaacca gttacaggct
421 gccagtggcc ccggcttcata tccggcttcgc gcctcgccgc ggcccaagcc ctgcctctc
481 ctggaaatag gtcaggatgt tagtccgggtt cgcgcgtgt ccactgcgc tgctccagct
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661 ggtcggtcgcc ccggccgggg agggctctgc agttgcgcag cttgctcccc ggccctttc
721 ccctccgctc cccggccctt cctgacgcgc ggcgtgacgt caccacgccc ggccggccgccc
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841 ccgagttactg cggtgagacg cagccggccca ggcgcggccgtt caacagccgc cagaagttaca
901 cgagggaaaccc ggcggccgggt gtgcgtgtgc gcccgtgtgc gggccggccgc gggggaggag
961 cgccggccgggg cggccggctgg gggccgggtgg catcatggac gagaagggtgt tcaccaaggaa
1021 gctggaccag tggatcgacg agctgaacga gtgcacgcag ctgtcccgat cccaggta
1081 gagcccttcgc gagaaggctt aagaaaatccct gacaaaaagaa tccaaacgtgc aagagggtcg
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1261 aggtatattat tcagttggaaa cagttacact gcttgcgtatctttaagggttc gttaccgtga
1321 acgcatcacc attcttcgag ggaatcatga gagcagacag atcacacacaag ttatgttt
1381 ctatgtgaa tggtttaagaa aatatggaa tgcaaatgtt tggaaatattt ttacagatct
1441 ttttgcgttactt cttctctca ctgccttgggtt ggatggccag atcttcgttc tacatgggtgg
1501 tctctcgcca tctatagata cactggatca tatcagagca cttgatcgcc tacaagaatgt
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1621 gggtatataat cctcgaggag ctgggttacac ctttggccaa gatatttgc agacattaa
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1741 ctgggtccat gaccggatgc tagtaacgtt tttcgtgtccatgacc tttatcggtt
1801 tggttaaccaa gctgcaatca tggaaatgttgcataacttca aaatacttctt tcttgcgtt
1861 tgaccggca cctcgtagag gggccacatggatgttgcgttccatgacc actacttctt
1921 gtaatggaaat tttaaaacttgc tagtatttgc ccatgaaatccatgcacc taatggaaat
1981 gggaaatggca acagtaacttcaaaatgttgcataaaatccatgcacc aaacttgcgtt
2041 tcacatggac caaaatgttgcataaaatccatgcacc ctcttgcgttccatgcacc
2101 gaccacttta gaatgaaatccatgcacc ttatgttgcgttccatgcacc acatgttgcgtt

PP2A Catalytic Subunit

2161 agtttctggc atagcgctat ttgttagttac ttttgcttcc tctgagagac tgcaagataat
2221 aagatgtaaa cattaacacc tcgtgaataac aatttaactt ccatttagct atagctttac
2281 tcagcatgac tgtagataag gatagcagca aacaatcatt ggagcttaat gaacatffff
2341 aaaaataatt accaaggcct cccttctact tgtgagttt gaaatgttc tttttatttt
2401 cagggataacc gtttaattta attatatgtt ttgtctgcac tcagtttattt ccctactcaa
2461 atctcagccc catgttggc tttgttattt tcagaacctg gtgagttt ttgaacagaa
2521 ctgttttttc ccottcctgt aagacgatgt gactgcacaa gagcaactgca gtgttttca
2581 taataaaactt gtgaactaag aactgagaag gtcaaatttt aattgtatca atgggcaaga
2641 ctgggtctgt ttattaaaaa agttaaatca attgagttttt ttttagaatt ttagacttg
2701 taggtaaaaat aaaaatcaag ggcactacat aacctcttg gtaactcctt gacattttc
2761 agattaactt caggatttat ttgtatttca catattacaa ttgtcacat tttgggtgtg
2821 cacttgtgg gttttccctg catattaact ttttgtaag aaaggaaatc tttgtgtgtt
2881 cagtaagact taattgtaaa accatataac ttgagattt agtcttggg ttgtgtttta
2941 ataaaacagc atgttttcag gtagag

PP2A PR55 Regulatory Subunit

LOCUS PP2APR55 2131 bp ss-mRNA PRI 09-MAY-1991
DEFINITION Human protein phosphatase 2A alpha subunit mRNA, complete cds.
ACCESSION M64929 J05328
KEYWORDS protein phosphatase-2A subunit-alpha; regulatory subunit.
SOURCE Human lung fibroblast cell line WI38, cDNA to mRNA.
REFERENCE 1 (bases 1 to 2131)
AUTHORS Mayer,R.E., Hendrix,P., Cron,P., Matthies,R., Stone,S.R., Goris,J.,
Merlevede,W., Hofsteenge,J. and Hemmings,B.A.
TITLE Structure of the 55 kDa regulatory subunit of protein phosphatase
2A: Evidence for a neuronal specific isoform
JOURNAL Biochemistry 30, 3589-3597 (1991)
COMMENT PP2A consists of regulatory subunits and one catalytic subunit.
In their review, "Protein Phosphatase 2A and the Regulation
of Human Papillomavirus Gene Activity, Schegget and Noordaa
(in Human Papillomaviruses, 121-9) report that PP2A is
inactivated by high expression of its regulatory subunit,
PR55Beta. This inhibition is linked to the activation of the LCR
region of HPV-16. Thus, the authors suggest that the regulation
of the HPV-16 promoter is correlated to cellular protein
phosphorylation.

NCBI gi: 190421
BASE COUNT 665 a 421 c 460 g 585 t
ORIGIN

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1 ccggccgcat ccggccctc taccggcca tccccaggta aggggggtga gttcaggaag
 61 cggagacccc gaggaaccca gcagggtcac cattgcacg gcaacatggc aggagctgga
121 ggagggaaatg atattcagtg gtgttttct caggtgaaag gagcagttaga tgatgtgta
181 gcagaagcag atataatttc tacagttagaa tttaatcatt ctggagaatt actagcaaca
241 ggagataaaag gtggtagagt tgtcatctt caacaggagc aggagaacaa aatccagtct
301 catagcagag gagaatataa tgtttacagc accttccaga gccatgaacc agagttgac
361 tacttggaaaa gtttagaaat agaagaaaag atcaataaaaa ttagttgggtt accccagaaa
421 aatgcgtcgtc agtttttatt gtctaccaat gataaaaacaa taaaattatg gaaaatcagt
481 gaaagggaca aaagaccaga agggtataac ttgaaagagg aggttggaaat gtatagagat
541 cctactacag ttactacact acgagtgcac gtcttaggc ctatggatct aatggttgag
601 gccagtcac gaagaatatt tgccaatgct catacatatc acatcaactc aatttctatt
661 aatagtgatt atgaaacata ttatctgca gatgatttgc ggattaatct ttggcatctg
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1201 ttcaaatgtt ttgacagaaaa cacaaagcga gacataaccc tagaagcattc gcgaaaaaac
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1321 gatgaaataaa gtgttgcacag cctagacttca aataagaaaa tccttcacac agctggcac
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1441 gtgaattagg ttggcatttc ctgcgcgttgc aacccacttc ctgccttagtt gagatagttg
1501 aatcttagcat tcgttcctat aaaagagaga ggtccattgt ggcccccctt tccagtgttt
1561 gacagtgtgc cattcgacaa cacattgtta tagctacatg gagaagctc tggatgtca
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2101 cgccgtgtct ggactgtaaa ataaggaaaa g
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PP2A PR65 Regulatory Subunit

LOCUS PP2APR65 2205 bp ss-mRNA PRI 22-OCT-1992
DEFINITION Human protein phosphatase 2A regulatory subunit alpha-isotype
(alpha-PR65) mRNA, complete cds.
ACCESSION J02902
KEYWORDS protein phosphatase-2A regulatory alpha-subunit.
SOURCE Human HeLa cell, cDNA to mRNA, clone lambda-HHPR65-3.
REFERENCE 1 (bases 1 to 2205)
AUTHORS Hemmings,B.A., Adams-Pearson,C., Maurer,F., Mueller,P., Goris,J.,
Merlevede,W., Hofsteenge,J. and Stone,S.R.
TITLE Alpha and beta-forms of the 65-kDa subunit of protein phosphatase
2A have a similar 39 amino acid repeating structure
JOURNAL Biochemistry 29, 3166-3173 (1990)
COMMENT PP2A consists of regulatory subunits and one catalytic subunit.
In their review, "Protein Phosphatase 2A and the Regulation
of Human Papillomavirus Gene Activity, Schegget and Noordaa
(in Human Papillomaviruses, 121-9) report that PP2A is
inactivated by high expression of its regulatory subunit,
PR55Beta. This inhibition is linked to the activation of the LCR
region of HPV-16. Thus, the authors suggest that the regulation
of the HPV-16 promoter is correlated to cellular protein
phosphorylation.

Draft entry and printed sequence [1] kindly submitted by
B.A.Hemmings, 23-MAR-1990, for release after publication.

NCBI gi: 189427

BASE COUNT 447 a 676 c 639 g 443 t
ORIGIN

1 gaattccgggt ttcacttgc gacgttgtcc agtccagca cttggcaac tccccagct
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121 tgacggaaaa gggacggagc caagatggcg gggccgacg ggcgcagtc gctgtacccc
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301 cttttcccta cagatccat ctatgatgaa gatgagggtcc ttctggccctt ggccagaacag
361 ctggaaacct tcactaccct ggtgggaggc ccagagtaacg tgcactgcctt gctgccaccg
421 ctggagtcgc tgccacagt gggaggagaca gtggtgccgg acaaggcagt ggagtccctt
481 cggccatctt cacacgagca ctggccctt gacctggagg cgcaacttggt gccgcttagtg
541 aagcggctgg cggccggcgta ctggttcacc tcccgacactt cggccctggg ccttccctcc
601 gtctgttacc ccacggatgtc cagtgctgtt aaggcggaaat ttgcacagta ctggccggaa
661 ctgtgttcgtt atgacaccccc catggtgccgg cggccggcgat cctccaaat gggggagtt
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1741 gttgcataatg tccgttcaa tttggccaaatg tttttgttgcata gataggggcc catcctggac
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1861 gttgcataatg tggccatccat gactacgttca tttttgttgcata gataggggcc catcctgg
1921 aagaggagca aacactggcc tctgggttcc accctccaaat cccaccaatg ccctctttgg
1981 gggacactg gggggccctt ggtgttgcactt ccctgttgcat ggtctgttgc caggccctt
2041 ccccaacatc ggttccctt cttcccaatc tggccatggc tggccatggc tttttgttgc
2101 cggcttgggg agtggccgggtt tggacaggac agtggccatggg ggttgcataatg
2161 catccatggaaatc agccatggag ccggaggtgg caatttcacccg aatcc

Transcriptional Enhancer Factor

LOCUS TEF 4443 bp ds-DNA PRI 28-JUL-1992
DEFINITION Transcriptional enhancer factor DNA, complete CDS.
ACCESSION M63896
KEYWORDS trans-acting transcriptional activator; transcription enhancer.
SOURCE Homo sapiens (library: ZAP-II random primed cDNA) DNA.
REFERENCE 1 (bases 1 to 4443)
AUTHORS Xiao,J.-H., Davidson,I., Matthes,H., Garnier,J.-M. and Chambon,P.
TITLE Cloning, expression, and transcriptional properties of the human
enhancer factor TEF-1
JOURNAL Cell 65, 551-568 (1991)
BASE COUNT 1226 a 1040 c 969 g 1208 t
ORIGIN

1 cgccggccgc cgccggcgcc caccaagcac ttgcacact cgcttccacc ctgcgggcca
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121 gccccccac ccctccaccc ttgcggcggc ggcgggcgc agcccagcgc gccagccggc
181 cccggggcag gagcgggtgc aaggcaggggt ggggtggccg gggccaggga cccggagccg
241 gggaggggcgc cgggcaccga gcagaggggcg ggggaagccg cggcgaagtt tgccctggac
301 tcgccccggc ctgcgggtgc tccctggcc gaggactgt gtcgcgtg cccggccgc
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1981 gaagtcattt tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg
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2521 tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg
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3001 aacttagggctt tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg tttttatgg
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Transcriptional Enhancer Factor

3121 atctcaatgt gtctgttagac ttagatacat cctttgaag cacatccatt tcttttagcgt
3181 ctctcagtaa gttacagtac ttgtttgact taggtttaag aggcccagct acctatctct
3241 gacctttca aataggctca ttggggagat tcttttgcctt ggagagattc aactttccaa
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4081 agccaggagg gctttgggat tttcttttccaaagcacaaa aatactgggacccaaagaaga
4141 acagcttagag gacaactctg ttggcacaga gacggggaca gcccagtctg ctgacccctac
4201 agggtcagtg ggcccccctg gtgcttcacc acctgcatttgc tcttgcctc aatgccttgc
4261 cagttgagtt ttctgggtt ctatgattga ctttgagtt tactccttgc tcttacaaca
4321 tttctaaagga tttttaaaag tttacttctt gtcttgcatttct tctaaagctt tctccaggac
4381 agatattttc cctgtcttaa ccactggtcc agtcatccca gtgggcttctt ctgtctct
4441 ccc

Transforming Growth Factor

LOCUS TGFA 867 bp ss-mRNA PRI 19-APR-1994
DEFINITION Human (cell line 1027 F57) transforming growth factor-alpha mRNA,
complete cds.
ACCESSION K03222
KEYWORDS growth factor; transforming gene; transforming growth factor-alpha.
SOURCE Human renal carcinoma cell line 1027 F57, cDNA to mRNA, clone
pTGF-Cl.
REFERENCE 1 (bases 1 to 867)
AUTHORS Derynck,R., Roberts,A.B., Winkler,M.E., Chen,E.Y. and Goeddel,D.V.
TITLE Human transforming growth factor-alpha: Precursor structure and
expression in *E. coli*
JOURNAL Cell 38, 287-297 (1984)
COMMENT Pietenpol et al. (Cell 6: 777-785) report that HPV-16 and HPV-18
infection inhibits growth suppression by TGF-B1 in
human foreskin keratinocytes. The authors suggest that the inhibitory
capability of TGF-B1 is linked to the down-regulation of c-myc and
that HPV-16 E7 expression interferes with this pathway.
[1] also sequenced exons 1 and 2 of this gene from the genomic DNA
library of Lawn et al. (see separate entries).
BASE COUNT 175 a 240 c 254 g 198 t
ORIGIN Chromosome 2p13
1 aggctggaga gcctgctgcc cgcccgcccc taaaatggtc ccctcggctg gacagctcgc
61 cctgttcgct ctgggttattg ttgtggctgc gtgccaggcc ttggagaaca gcacgtcccc
121 gctgagtgc aaccgcggcc tggctgcago agtggtgtcc cattttaatg actgcccaga
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241 agcatgtgc tgccattctg ggtacgttgg tgcacgtgtt gagcatgcgg acctcctggc
301 cgtggggct gccaGCCaga agaAGCAGGC catCACCGCC ttgggtgtgg tctccatcgt
361 ggccctggct gtccttatca tcacatgtgt gctgatacac tgctGCCagg tccgaaaaca
421 ctgtgagtgg tgccggggcc tcatctgcgg gcacgagaag cccagcggcc tcctgaaggg
481 aagaaccgtc tgctgccact cagaaacagt ggtctgaaga gcccagagga ggagtttggc
541 cagggtggact gtggcagatc aataaagaaa ggcttcttca ggacagcact gccagagatg
601 cctggggctg ccacagacct tcctacttgg cctgtaatca cctgtgcagc cttttgtggg
661 ccttcaaaac tctgtcaaga actccgtcgg ctgggggtt ttcagtgtga cctagagaag
721 aaatcagcgg accacgattt caagacttgt taaaaaaagaa ctgcaaagag acggactct
781 gttcacctag gtgagggttg tgcaGAGTT ggtgtctgag tccacatgtg tgcaGTTGTC
841 ttctggccagc catggattcc aggccgt

Tumor Necrosis Factor

LOCUS TNF 7112 bp ds-DNA PRI 15-JUN-1989
DEFINITION Human tumor necrosis factor and lymphotoxin genes, complete cds.
ACCESSION M16441
KEYWORDS lymphotoxin; tumor necrosis factor.
SOURCE Human placenta DNA, clone pTNF186.
REFERENCE 1 (bases 1 to 7112)
AUTHORS Nedospasov,S.A., Shakhov,A.N., Turetskaya,R.L., Mett,V.A.,
Azizov,M.M., Georgiev,G.P., Korobko,V.G., Dobrynin,V.N.,
Filippov,S.A., Bystrov,N.S., Boldyreva,E.F., Chuvpilo,S.A.,
Chumakov,A.M., Shingarova,L.N. and Ovchinnikov,Y.A.
TITLE Tandem arrangement of genes coding for tumor necrosis factor
(TNF-alpha) and lymphotoxin (TNF-beta) in the human genome
JOURNAL Cold Spring Harb. Symp. Quant. Biol. 51, 611-624 (1986)
COMMENT Kyo et al. (Virology 200: 130-9) demonstrated that tumor necrosis
factor (TNF) alpha represses transcription of the early HPV-16
genes through responsive elements in the LCR. Malejczyk et al.
(Int J Cancer 56: 593-8) reported this inhibition for weakly
tumorigenic cell lines, but not for highly tumorigenic cell lines.
Further Malejczyk et al. demonstrated that TNF-alpha receptor
expression is reduced in the higly tumorigenic lines and that this
reduction may inhibit the autocrine TNF-alpha-mediated growth
limitation.

Draft entry and computer-readable sequence for [1] kindly submitted
by C.V.Jongeneel, 02-OCT-1988.

BASE COUNT 1676 a 2005 c 1865 g 1566 t
ORIGIN 1 bp upstream of EcoRI site; chromosome 6p21.3.

1 gaattctcgaaacttccttttgtaaaaactttggaaagggtgtctccacattgatcctggaa
61 atgtgtgttttatttgggggttatataaatctttctgtggaaagccacctgaaatgcaggaa
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Tumor Necrosis Factor

6121 tattaccccc tccttcagac accctaacc ttttctggct caaaaagaga attggggct
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7021 cccaaacagaa tattccccat cccccaggaa acaagagctt gaacctaatt acctctccct
7081 cagggcatgg gaatttccaa ctctggaaat tc

LOCUS YY1 2353 bp ss-mRNA PRI 25-MAY-1993
 DEFINITION Homo sapiens GLI-Krupple related protein (YY1) mRNA, complete cds.
 ACCESSION M77698
 KEYWORDS GLI-Krupple related protein.
 SOURCE Homo sapiens cDNA to mRNA.
 REFERENCE 1 (bases 1 to 2353)
 AUTHORS Shi,Y., Seto,E., Chang,L.-S. and Shenk,T.
 TITLE Transcriptional repression by YY1, a human GLI-Krupple related protein, and relief of repression by adenovirus E1a protein
 JOURNAL Cell 67, 377-388 (1991)
 COMMENT YY1 has been shown to be involved in transcriptional repression of both HPV-16 and HPV-18 (May,M., Dong,X.-P., Beyer-Finkler,E., Stubenrauch,F., Fuchs,P., and Pfister,H., EMBO J.:13, 1460-1466 (1994); Bauknecht,T., Angel,P., Royer,H.-D., and zur Hausen,H., EMBO J.:11, 4607-4617 (1992)). It is thought to interfere with transcriptional initiation by binding to sites contained within a transcriptional silencer domain in the LCRs of these HPV types. In HPV-16, this silencer domain is contained within a naturally occurring deletion mutation which was found by May et al. to be present in 40% of the HPV-16 DNA in a lymph node metastasis of a cervical cancer. May et al. contend that this deletion may represent a means by which expression of oncogenic genes is de-regulated in episomal copies of HPV-16 DNA. Since at least one-third of HPV-16-positive cervical cancers contain only extra-chromosomal viral DNA, this deletion may be an important alternative to the destruction of the E2 reading frame, and thereby the E2-dependent negative regulation of E6/E7 expression, that occurs during integration of the viral DNA into the host genome. Mutations of YY-1 binding sites in HPV-18 DNA has been shown to affect transcriptional activity differently in malignant and non-malignant cell lines. In non-malignant cells, YY1 binding-site mutations led to increased transcriptional activity, whereas in certain malignant cell lines, transcriptional activity was apparently reduced by the mutations. This may point to more complex interactions between YY1 and other cellular factors.

BASE COUNT 625 a 598 c 608 g 522 t

ORIGIN

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1 cgccgagacg agcagcgccc gagcgagcgc gggcgccggc gcaccgaggc gagggaggcg
61 gggaaagcccc gcccggccg ccccgccccg cccttccccc gccgccccgc ccctctcccc
121 cccggccgtc gccgccttc tccctctgc ttcccttcccc acggccggcc gcctcctcgc
181 cccggccggcc gcagccgagg agccgaggcc gccgccccgg tggcgccgga gcctcagcc
241 atggcctcg ggcacaccc ctacatcgcc acggacggct cggagatgcc ggcggagatc
301 gtggagctgc acgagatcga gttggagacc atcccggtgg agaccatcga gaccacagtg
361 gtggggcgagg aggaggagga ggacgacgcgacgaggacg gcccgggtgg cgaccacggc
421 ggcggggcg gccacgggca cgccggccac caccaccacc accatcacca ccaccaccac
481 cccggccatga tcgctctgca gccgctggtc accgacgcacc cgaccagggt gcaccaccac
541 caggaggtga tcctggtgc gacgcgcgag gaggtgggtgg gcccggacgc ctcggacggg
601 ctgcgcgcgc aggacggctt cgaggatcatcg attctcatcc cggtgccgc gccggccggc
661 ggcgacgcgact acacattga acaaacgcgt gtcaccgtgg cggcgccgg caagagcgcc
721 ggcggccggct cgtcgtcgct gggaggccgc cgcgtcaaga agggcgccgg caagaagagc
781 ggcaagaaga gttacctcg cggcgccggc ggcgcggccg gcccggccgg cgccgacccg
841 ggcaacaaga agtggggagca gaagcgaggta cagatcaaga ccctggaggg cgagttctcg
901 gtcaccatgt ggtcctcaga tgaaaaaaaaa gatattgacc atgagacagt ggttgaagaa
961 cagatcattg gagagaacttcc acctccgtat tattcagaat atatgacagg aaagaaactt
1021 cctccctgg gaataccctgg cattgaccc tcagatccca aacaactggc agaatggct
1081 agaatgaaacg caaaaaat taaaagaaat gatgtccaa gaacaatagc ttgcctcat
1141 aaaggctgca caaagatgtt cagggttccaa tcggccatga gaaaacatct gcacacccac
1201 ggtcccgag tccacgtctg tgcagaatgt ggcaaaagggt ttgtttagag ttcaaaaacta
1261 aaacgacacc aactgggtca tactggagag aagcccttc agtgcacgtt cgaaggctgt
1321 gggaaacgtt ttcactggta cttcaatttgc cgcacacatg tgcaatcca tacccggagac
1381 aggcctatg tggccctt cgtatgggtgt aataagaatgt ttgctcgtc aactaacatg
1441 aaatctcaca tcttaacaca tgctaaggcc aaaaacaacc agtggaaaaga agagagaaga
1501 cccttctcgcc acacggaaat catcttccag aagtgttattt gggataaat atgcctctcc
1561 tttgttatatttattttagga agaattttaa aaatgaaatcc tacacaccta agggacatgt

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YY1

1621 tttgataaaag tagtaaaaat taaaaaaaaaa aaactttact aagatgacat tgctaagatg
1681 ctctatcttg ctctgtatc tcgttcaaa aacacagtgt tttgtaaag tgtggtccca
1741 acaggaggac aattcatgaa cttcgcatca aaagacaatt ctttatacaa cagtgcataaa
1801 aatgggactt ctttcacat tcttataaaat atgaagctca cctgttgctt acaattttt
1861 taattttgt a tttccaagt gtgcatttg tacactttt tgggatatg ctttagtaatg
1921 ctacgtgtga ttttctgga ggttgataac ttgcctgca gtagatttc tttaaaagaa
1981 tgggcagttt catgcatact taaaagttt tttcctgtaa aaaaaaaaaa agtttatata
2041 gttttgtttt cttatcttataat ttgggttgc ttctttgatg ttaacacatt ttgtataatt
2101 gtatcgata gctgtattga atcatgtatg atcaaatattt agatgtgatt taatagtgtt
2161 aatcaattta aaccattttt agtcactttt ttttccaaa aaaatactgc cagatgctga
2221 tgttcagtgt aatttctttg cctgttcagt tacagaaagt ggtgctcagt tgtagaatgt
2281 attgtacctt ttaaacacctg atgtgtacat cccatgtaac agaaaggca acaataaaaat
2341 agcaatccta aag

LOCUS c-myc 8082 bp ds-DNA
 DEFINITION Human (Lawn) c-myc proto-oncogene, complete coding sequence and flanks.
 ACCESSION J00120
 KEYWORDS Alu repeat; c-myc proto-oncogene; myc oncogene; proto-oncogene; repeat region; transforming gene.
 SOURCE Human DNA (genomic library of Lawn et al.), clones lambda-M1 [1], and pUC9-myc [2].
 REFERENCE 1 (bases 3507 to 7559)
 AUTHORS Colby,W.W., Chen,E.Y., Smith,D.H. and Levinson,A.D.
 TITLE Identification and nucleotide sequence of a human locus homologous to the v-myc oncogene of avian myelocytomatosis virus MC29
 JOURNAL Nature 301, 722-725 (1983)
 REFERENCE 2 (bases 1 to 8082)
 AUTHORS Gazin,C., Dupont,S., de Dinechin,D., Hampe,A., Masson,J.M., Martin,P., Stehelin,D. and Galibert,F.
 TITLE Nucleotide sequence of the human c-myc locus: provocative open reading frame within the first exon
 JOURNAL EMBO J. 3, 383-387 (1984)
 COMMENT In several cases of HPV-positive cancer, Couturier et al. (J Virol 65: 4534-8) determined that HPV sequences integrate near the myc locus. In several instances, this integration disrupted the integrity of the proto-oncogene sequence or its regulation.

The myc gene is the cellular homologue of the transforming gene carried by the avian myelocytomatosis virus MC29. Unlike the ras proto-oncogenes which obtain transforming potential through mutations within their coding exons (namely mutations within codon 12), the myc gene identified as the cause of Burkitt lymphomas acquires its transforming potential through defects of either transcriptional or translational control. Thus it is not an altered gene product that induces tumors, but a normal product that is present either in the wrong quantity or at the wrong time in the life cycle of the cell.

[2] notes an open reading frame upstream of the c-myc coding exons with an 'atg' start codon at bases 2304-2306 and a 'tag' stop codon at bases 2868-2870. However other researchers have used c-myc and v-myc DNA sequences to probe for mRNA's with homology to c-myc in various human cell lines and none of them have noted any mRNA's beginning upstream of bp 2328 (see other human c-myc entries).

The t(8;14) translocation site in the Burkitt lymphoma cell line BL22 occurs between bp 1316 and 1317 of this sequence.

BASE COUNT 1850 a 2115 c 2135 g 1982 t
 ORIGIN 198 bp upstream of Sau96A site, on chromosome 8 (q24).
 1 agcttggtttgcgcgttttag gtttttttttcgttatgt acttgtaat
 61 tatttcacgt ttggccattac cggttctcca tagggtgatg ttcattagca gtggtgatag
 121 gttaattttc accatctttt atgcgggttga atagtcacct ctgaaccact ttttcctcca
 181 gtaactccctc ttcttcggaa cttctgtcgag ccaacctgaa agaataacaa ggaggtggct
 241 ggaaacttgt tttaaggaaac ccgcgttgcct tcccccgctg gaaacccgtc acctcgacg
 301 ctccgtctcc tgccccccacc tgaccccccgc cctcgttgc atccaggcgc gatgatctt
 361 gctgccagta gagggcacac ttactttact ttgccaaacc tgaacgcggg tgctgcccag
 421 agagggggcg gagggaaaaga cgctttgcag caaaatccag catagcgatt gttgttccc
 481 cgcgttgcg gcaaaaggct ggaggcagga gtaatttgca atccattaaag ctgaatttgt
 541 cagtgcacgt gatttggaaat ctactatatt cacttaaacat ttgaacgcgt agctgcaaac
 601 tcaacgggttataaaccatc ttgaacagcg tacatgtcat acacacaccc ctcccccccg
 661 aatttttttc tctttttggat gtgggtggagg gagagaaaaat ttacttaaa atgcctttgg
 721 gtgaggggacc aaggatgaga agaatgtttt ttgttttca tgccgtggaa taacacaaaa
 781 taaaaaaatcc cgagggaaatacattat attaaataat gatcattca gggagcaaac
 841 aaatcatgtg tggggctggg caactagctg agtgcgaagcg taaataaaat gtgaatacac
 901 gtttgcgggt tacatacagt gcactttcac tagtattcgt aaaaaattgt gatgttgc
 961 acttagaaaaat taatgcctgg aaggcagcca aattttaaat agtcaagac tcccccccc
 1021 ccccaaaaaa aggcacggaa gtaatactcc tctcccttcc tttgtatcaga atcgatgtat

1081 tttttgtgca tgaccgcatt tccaataata aaagggaaa gaggacctgg aaaggaatta
 1141 aacgtccgtt ttgtccgggg agggaaagagt taacggtttt ttcacaagg gtctctgctg
 1201 actccccccg ctcggcac aagctctcca ctggccctt ttaggaagtc cggtcccgcg
 1261 gttcgggtac ccctgcgccccc tcccatattc tcccgtag cacctttagt ttctccaaa
 1321 cccggcagcc cgagactgtt gcaaaccggc gccacaggc gcaaaggaa tttgtcttctt
 1381 ctgaaacctg gctgagaaat tgggaactcc gtgtgggagg cgtgggggt ggacgggtggg
 1441 gtacagactg gcagagagca gcaacactcc ctctcgccct agcccagtc tggAACAGGC
 1501 agacacatct caggctaaa cagacgcctc cgcacgggg cccacggaa gcctgagcag
 1561 gcggggcagg agggcggta tctgctgctt tggcagcaaa ttggggact cagtcgtggg
 1621 ggaagggttcaatccatgat agctgtgcat acataatgca taatacatgta ctccccccaa
 1681 caaatgcaat gggagtttat tcataacgcg ctctccaagt atacgtggca atgcgttgct
 1741 gggattttt aatcattcta ggcattcggtt tccctctatgc ttcctcccta
 1801 tctacactaa catccacgc tctgaacgcg cgcattaa tcccttctt tccctccactc
 1861 tccctggac tcttgcataa agcggggccc ttcccccagg cttagggagg cgccctgcag
 1921 cctggatcgcg gctggcgtg ggggtggcg cgcaatgcgt tctctgtgtt gaggcagct
 1981 gttccgcctg cgatgattta tactcacagg acaaggatgc gtttgtcaa acagtaactgc
 2041 tacggaggag cagcagagaa agggagaggg ttgagaggg agcaaaagaa aatggtaggc
 2101 ggcgttagtt aattcatgcg gtcctttaatc tctgttaca tcctagagct agagtgcgt
 2161 gtcggccggc tgagtctctt cccacacttc cccacccccc ccacccccc cataagcgcc
 2221 cttcccggtt tcccaaagca gaggcggtgg gggaaaagaa aaaagatctt ctctcgctaa
 2281 tctccggcca cggccctttt ataatgcgag ggtctggacg gtcgaggacc cccgagctgt
 2341 gtcgtcgccg gcccacccg cggggccccc ggcgtccctg gtcctccccc tgcgtcgaga
 2401 agggcaggcgtt ttctcagagg cttggggggaaa aaaaacgg agggagggat cgcgctgagt
 2461 ataaaaagccg gtttccgggg ctttatctaa ctgcgtgttag taattccagc gagaggcaga
 2521 gggagcgagc gggccggccgg ctgggtggaa agagccggc ggcagagct ggcgtcgaaa
 2581 cgtctggaa agggagatcc ggacgaaataa gggggcttcg cctctggccc agccctcccg
 2641 ctgatcccc acccagcggtt ccgcacccct tgcgcacatcc acgaaacttt gcocatagca
 2701 gccccggcgc actttgcact ggaacttaca acaccccgac aaggacgcga ctctccgcac
 2761 gccccggggc tattctgccc atttggggac acttcccgcc cgctgccagg accccgttct
 2821 ctgaaaggct ctccctgcag ctgcattagac gtcgttattt ttccgggttag tggaaaacca
 2881 ggtaagcacc gaagtccact tgccttttaa ttatattttt tatcacttta atgtcgat
 2941 gagtcgaatg cttaaatagg gtgtcttttccatccctt ggcatttttgcacttttcc
 3001 agagtagtta tgtaactgg ggtctggggggggggggtaatc cagaactggaa tggggtaaaa
 3061 gtgacttgc aagatggggagg aggaaaggc agagggaaaa cggggatggg ttttaagact
 3121 acccttcga gattctgcctt ttagatattt attcacgcgtt actcccgcc ggtcgacat
 3181 tctgttttaat tgcgtctgg gttttgggggggggggggggggggggggggggggggggggg
 3241 gggcggaaatcccttcgcatttgcgttccggatgggggggggggggggggggggggggggggg
 3301 agccagatcg ctcccgaccc gtcgttttttcccttcccttcccttcccttcccttcccttcc
 3361 gtcaccgcata ttttcgcacccgg
 3421 cgcggcgatttccacccgccttgcatt
 3481 caataataca att
 3541 caggcaggggaaaagg
 3601 aaaccaggcgcgaatctccgc acccagcccttgcgttgcgttgcgttgcgttgcgttgc
 3661 cctcgccccc gagatgcggg ggaactgcga ggagcgggggggggggggggggggggggggg
 3721 ctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgtt
 3781 gcatctccgttatttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgc
 3841 ttatgg
 3901 agaaaaagctg gaaaaaggag tgg
 3961 tggaggg
 4021 cgagttggaa cagccgcacccgg
 4081 ggcggccaggcgcgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgc
 4141 ggcaattgttcccttcacc ggcacccccc ggggggggggggggggggggggggggggggg
 4201 cgatttcttcgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgtt
 4261 cggggggctcg gggggccacca acccgctgggttgcgttgcgttgcgttgcgttgcgtt
 4321 actgtccaaatgggggtggaaa ggggtggcccttatttttttttttttttttttttttttt
 4381 ttatggggat agtctgcgg
 4441 ccagggtttccgcaccaaggccctttaacttcaagactgcgttgcgttgcgttgcgttgc
 4501 tccagcgcacccgg
 4561 tccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgc
 4621 agcaggcgcacccgg
 4681 agtctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgtt
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 4861 gtttcatctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgtt

4921 tgtggagcggtttctcgccgcccggccaaaggcttcaga gaagctggcc tcctaccagg
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 6541 tccagagacc ttt
 6601 att
 6661 acaagaagat gagaagaaa tggatgttttttttttttttttttttttt
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 7381 aaagaacttttttttttttttttttttttttttttttttttt
 7441 tgtt
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 7561 ccttagtatttttttttttttttttttttttttttttttt
 7621 aagttgttttttttttttttttttttttttttttttttt
 7681 gccaaatcttggatgttttttttttttttttttttttttt
 7741 cctgttttttttttttttttttttttttttttttttttt
 7801 ctctgttggatgttttttttttttttttttttttttttt
 7861 ttccctggatgttttttttttttttttttttttttttt
 7921 gagaaggatgttttttttttttttttttttttttttt
 7981 aaggccatataatgttttttttttttttttttttttt
 8041 ttaaccaggatgttttttttttttttttttttttttt

Cyclin A

LOCUS cyclinA 1181 bp ds-DNA PRI 21-JUN-1993
DEFINITION Human cyclin A (CCNA) gene sequence.
ACCESSION M96390
KEYWORDS cyclin A.
SOURCE Homo sapiens DNA.
REFERENCE 1 (bases 1 to 1181)
AUTHORS Nikaido,T. and Yamamoto,M.
TITLE p34cdc2 and cyclin A expression may be suppressed by RB and/or p53
JOURNAL Unpublished (1992)
COMMENT In the S-phase of the cell cycle, cyclin A complexes with E2F (a cellular transcription factor), p107, and cdk2 kinase. Arroyo et al. (Mol. Cell Biol. 13: 6537-46) report that the E7 protein of HPV-16 associates with the E2F-cyclin A complex. Arroyo et al. (Mol. Cell Biol. 13: 6537-46) further note that the high-risk HPV has a higher binding efficiency to the E2F-cyclin A complex than the low-risk HPV type. Steinmann et al. (Oncogene 9: 387-94) report a 20-fold increase in the level of cyclinA as a result of HPV-16 infection in human keratinocytes. Pagano et al. (Oncogene 7: 1681-6) report that the E2F-cyclin A complex can occur in HPV-18 infected cells independent of pRB.
BASE COUNT 302 a 279 c 262 g 338 t
ORIGIN
1 aagctttgtatattttata ttatatatata aatataaaaa ttgtttaaag gcacgtatag
61 ttaagagagt ttatattttaa taaggtcata ttgtttttac tatgtttaaa aaactttact
121 ctgaaaggaa cataattttata tcttagtcac tagaacgtca ttgtgtttt tgggttgc
181 acagcttggg gaaaaattaga aaaaattaa tgactgattt gaatatttg taatgcactg
241 ctattttata tatatatcaa ctagttca aggtgcattc taaaattaaat tgcatcttca
301 ttagaaaaaa taaaagcat aaaacacaat ttctggttac tatgaataaa cgctaaatg
361 ttaagatgac attacagtct tgacacttga gtactgtatt actatgtgag ctccgtgtta
421 aataatttat gcacattttataatcctaa aaccatatga ctgttagttat tagccctat
481 taacacataa gaaaacggag aatcgagat actgaaaaac gtccccaga ttttagacct
541 ttggaaaaag tcacttaagc taactagacg tcccagagct aaaggctggg caacccaaat
601 gatagtgcgc aaagtttaat tccgttaat tccctaaag gcttagagtc agttcggac
661 agcctcgctc acttagtggc tcaactttaaa ataatcgaa gggtcgggcc ctaaatccta
721 cctctccccc ccccgcgca ggttttctc ccccccagc cagttgttt ctccctctg
781 ccccgccccct gtcagtttc cttggttta cccttcactc gcccgtaccc tggcccttg
841 aatgacgtca aggccgcgag cgctttcatt ggtccatttc aatagtcgcg ggataacttga
901 actgcaagaa cagccgcccgc tccagcgggc tgctcgctgc atctctggc gtcttggct
961 cggcacgtg ggcaacttgcct gcctgcgcct ttgcacacct cctcgccct gggtggctc
1021 gagctgggtg agcgagcggg cggctggta ggctggctg ggctgcgacc ggoggctacg
1081 actattctt ggccgggtcg gtgcgagtgg tcggctggc agagtgcacg ctgtttggcg
1141 ccgcaggctg atcccgccgt ccactcccg gaggcgtat g

LOCUS GCORTICOIDR 2518 bp ds-DNA PRI 15-JUN-1990
 DEFINITION Human glucocorticoid receptor gene, partial.
 ACCESSION M32284
 KEYWORDS glucocorticoid receptor.
 SOURCE Human blood leukocyte DNA.
 REFERENCE 1 (bases 1 to 2518)
 AUTHORS Zong,J., Ashraf,J. and Thompson,E.B.
 TITLE The promoter and first, untranslated exon of the human glucocorticoid receptor gene are GC rich but lack consensus glucocorticoid receptor element sites
 JOURNAL Mol. Cell. Biol. 10, 5580-5585 (1990)
 COMMENT Authorin copy of sequence [Unpublished (1990) Univ. of TX Med. Branch, Galveston TX 77550] kindly submitted by B.E.Thompson, 22-FEB-1990.
 Mittal et al. (J Virol 67: 5656-9) identified three glucocorticoid response elements in the LCR of HPV-16 at position 7640, 7385, and 7474. This element, which binds the glucocorticoid receptor, is a partial palindrome and consists of the consensus TGTACANNNTGTCAT (Chan et al. J Virol 63: 3261-9). In the presence of glucocorticoids, up-regulation of the P\$_97\$ promoter is observed.

BASE COUNT 403 a 802 c 882 g 431 t
 ORIGIN

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  2461 ggcgcgtctg aattttactc gcccgaatat ttcaaccacc cccgcggccgc gcgagcc
  
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junB

LOCUS junB 2136 bp ds-DNA PRI 30-MAR-1994
DEFINITION Human transactivator (jun-B) gene, complete cds.
ACCESSION M29039
KEYWORDS c-myc proto-oncogene; transactivator.
SOURCE Homo sapiens DNA.
REFERENCE 1 (bases 1 to 2136)
AUTHORS Schuette,J., Viallet,J., Nau,M., Segal,S., Fedorko,J. and Minna,J.
TITLE jun-B inhibits the transforming and transactivating activities of c-jun
JOURNAL Cell 59, 987-997 (1989)
COMMENT Authorin copy of sequence [1] kindly submitted by J.D.Minna
10-OCT-1989.

Thierry et al. (J Virol 66: 3740-8) demonstrated that JunB binds AP-1 target sites within the LCR of the HPV-18 P\$_105\$ promoter during human keratinocyte infection. Mutation of these two target sites destroyed P\$_105\$ promoter activity in these cells. Bossy-Wetzel et al. (Genes Dev 6: 2340-51) reported that junB or c-jun are overexpressed during the aggressive fibromatoses or fibrosarcoma stage of BPV infection. They also correlated this overexpression to changes in fibroblast cell shape and anchorage dependence.

BASE COUNT 387 a 716 c 671 g 362 t
ORIGIN

1 cgcgagccgc ctcctccccct tccccacgct cgaggagggg ggccgcggggg ccoggctccg
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2101 cagcccgaggaa gtcggccaccctt tcccgctcc cccat

LOCUS p107Rb 3960 bp ss-mRNA PRI 06-MAY-1993
 DEFINITION Human retinoblastoma related protein (p107) mRNA, complete cds.
 ACCESSION L14812
 KEYWORDS cell cycle regulation protein; retinoblastoma protein;
 tumor suppressor.
 SOURCE Homo sapiens cDNA to mRNA.
 REFERENCE 1 (sites)
 AUTHORS Ewen,M.E., Xing,Y., Bentley-Lawrence,J. and Livingston,D.M.
 TITLE Molecular cloning, chromosome mapping and expression of the cDNA
 for p107 a retinoblastoma gene product-related protein
 JOURNAL Cell 66, 1155-1164 (1991)
 REFERENCE 2 (bases 1 to 3960)
 AUTHORS Zhu,L., van den Heuvel,S., Helin,K., Fattaey,A., Ewen,M.,
 Livingston,D., Dyson,N. and Harlow,E.
 TITLE Inhibition of cell proliferation by p107, a relative of the
 retinoblastoma protein
 JOURNAL Genes Dev. (1993) In press
 COMMENT p107, an Rb related protein, has been shown to complex with
 Rb (Carlotti et al. J Gen Virol 74: 2479-86). Davies et al.
 (J Virol 67: 2521-8) demonstrated that the HPV-16 E7 protein
 binds p107 and that this binding involves the Rb target site.
 B-myb, a gene involved in cell cycle progression, is
 inappropriately transcribed during G1 in HPV-16 E7 transactivated
 cells. Lam et al (EMBO J 13: 871-8) demonstrated that B-myb
 transcription is regulated through interactions with p107.

BASE COUNT 1257 a 771 c 869 g 1063 t
 ORIGIN

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Retinoblastoma-related p107

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